

—87. The method of claim 82, wherein the cell is a mouse embryonic stem cell.—

Sub 53 —88. A method of producing a transgenic embryo, comprising introducing a satellite artificial chromosome into an embryo.—

—89. The method of claim 88, wherein the embryo is a mouse embryo.—

—90. A method of producing a transgenic animal, comprising:
introducing a cell comprising an artificial chromosome comprising a heterologous nucleic acid into a female animal; and
allowing the cell to develop into a transgenic animal comprising an artificial chromosome comprising a heterologous nucleic acid.—

—91. The method of claim 32, wherein the transgenic animal is a mouse.—

Sub 105 —92. The method of claim 32, wherein the cell is a mammalian cell.—

B —93. A method of producing a transgenic animal, comprising:
introducing nucleic acid comprising a selectable marker into a first cell;
growing the cell under conditions that selectively permit the growth of cells containing the nucleic acid;
selecting a cell comprising a satellite artificial chromosome;
transferring the satellite artificial chromosome into a second cell, wherein the second cell is an animal cell;
introducing the second cell comprising the satellite artificial chromosome into a female animal; and
allowing the cell to develop into a transgenic animal comprising a satellite artificial chromosome.—

—94. The method of claim 93, wherein the satellite artificial chromosome is isolated prior to transferring it into a second cell.—

Sub 106 —95. A method of producing a transgenic animal, comprising:
introducing nucleic acid comprising a selectable marker into a first cell;

growing the cell under conditions that selectively permit the growth of cells containing the nucleic acid;

selecting a cell comprising a dicentric chromosome that comprises a *de novo* centromere;

growing the cell under conditions whereby a satellite artificial chromosome is produced;

transferring the satellite artificial chromosome into a second cell, wherein the second cell is an animal cell;

introducing the second cell comprising the satellite artificial chromosome into a female animal; and

allowing the cell to develop into a transgenic animal comprising a satellite artificial chromosome. —

—96. A method of producing a transgenic animal, comprising:

introducing nucleic acid comprising a selectable marker into a first cell;

growing the cell under conditions that selectively permit the growth of cells containing the nucleic acid;

selecting a cell comprising an artificial chromosome that comprises more heterochromatic nucleic acid than euchromatic nucleic acid;

transferring the artificial chromosome into a second cell, wherein the second cell is an animal cell;

introducing the second cell comprising the artificial chromosome into a female animal; and

allowing the cell to develop into a transgenic animal comprising an artificial chromosome that comprises more heterochromatic than euchromatic nucleic acid. —

—97. A method for producing a transgenic animal, comprising:

introducing an embryo comprising a satellite artificial chromosome into a female animal; and

allowing the embryo to develop into a transgenic animal comprising a satellite artificial chromosome. —

—98. A method for producing a transgenic animal, comprising:
introducing a fertilized oocyte comprising a satellite artificial chromosome into a female animal; and

allowing the embryo to develop into a transgenic animal comprising a satellite artificial chromosome. —

—99. A method for producing a transgenic animal, comprising:
introducing a mouse embryonic stem cell comprising a satellite artificial chromosome into an embryo;

introducing the embryo into a female animal; and

allowing the embryo to develop into a transgenic animal comprising a satellite artificial chromosome. —

—100. The method of claim 32, wherein the cell is a mouse cell. —

—101. A non-human transgenic embryo comprising a satellite artificial chromosome. —

—102. A non-human transgenic embryo comprising an artificial chromosome that contains more heterochromatic nucleic acid than euchromatic nucleic acid. —

—103. A non-human transgenic embryo comprising a satellite artificial chromosome, wherein the satellite artificial chromosome is obtained by a process comprising:

introducing nucleic acid comprising a selectable marker into a cell;
growing the cell under conditions that selectively permit the growth of cells containing the nucleic acid; and

selecting a cell that comprises a satellite artificial chromosome. —

—104. A non-human transgenic embryo comprising a satellite artificial chromosome, wherein the satellite artificial chromosome is obtained by a process comprising:

introducing nucleic acid comprising a selectable marker into a cell;
growing the cell under conditions that selectively permit the growth of
cells containing the nucleic acid;

selecting a cell comprising a dicentric chromosome that comprises a *de novo* centromere;

growing the cell under conditions whereby a satellite artificial
chromosome is produced; and

selecting a cell that comprises a satellite artificial chromosome. —

B' — 105. A non-human transgenic embryo comprising an artificial
chromosome, wherein the artificial chromosome is obtained by a process
comprising:

introducing nucleic acid comprising a selectable marker into a cell;
growing the cell under conditions that selectively permit the growth of
cells containing the nucleic acid; and

selecting a cell that comprises a minichromosome that comprises a
neocentromere, the nucleic acid and euchromatin. —

Please amend claims 32-34, 36-39, 41, 43, 44, 65, 67, 73 and 74 as follows:

sub D' 32. (Twice Amended) A method for producing a transgenic animal,
comprising introducing a cell comprising a satellite artificial chromosome into [an
embryonic cell] a female animal; and

allowing the cell to develop into a transgenic animal comprising a satellite
artificial chromosome

B² [and exposing the cell containing the satellite artificial chromosome to conditions
whereby a transgenic animal develops therefrom].

33. (Amended) The method of claim 32, wherein the [embryonic] cell is
a stem cell.

34. (Amended) The method of claim 32, wherein the [embryonic] cell is
in an embryo.

36. (Twice Amended) The method of claim [35]32, wherein [the product is the cystic fibrosis transmembrane regulatory protein, an anti-HIV ribozyme, or a tumor suppressor gene] the cell is an oocyte.

37. (Twice Amended) The method of claim [36]32, wherein the [anti-HIV ribozyme is an anti-*gag* ribozyme, and the tumor suppressor gene is p53] cell is a germline cell.

B³
38. (Twice Amended) The method of claim [35]32, wherein the [product comprises an antigen that upon expression induces a immunoprotective response against a pathogen in the transgenic animal] cell contains the satellite artificial chromosome in a pronucleus.

39. (Twice Amended) The method of claim [35]32, wherein the [product comprises a plurality of antigens that upon expression induce an immunoprotective response against a plurality of pathogens] cell is a zygote.

B⁴
41. (Twice Amended) The method of claim 32, wherein the [satellite artificial chromosome is introduced by cell fusion, microinjection, microcell fusion, electroporation, microprojectile bombardment or direct DNA transfer] the transgenic animal is a mammal.

sub D²
43. (Twice Amended) A method of producing a transgenic animal, comprising:

introducing [DNA] nucleic acid into a first cell;

growing the cell under conditions that selectively permit the growth of [a cell] cells containing the [DNA] nucleic acid;

B⁵
selecting a cell that comprises a minichromosome that is about 10 Mb to about 50 Mb that comprises a neo-centromere, the [DNA] nucleic acid and euchromatin;

transferring the minichromosome into a second cell, wherein the second cell is an animal cell; [and

exposing the animal cell containing the minichromosome to conditions whereby a transgenic animal develops therefrom]

introducing the cell comprising the minichromosome into a female animal;
and

allowing the cell introduced into the female animal to develop into a
transgenic animal comprising a minichromosome; wherein,

the [DNA] nucleic acid comprises DNA encoding a selectable marker and
a gene product or products;and

the DNA encoding the selectable marker and the DNA encoding the gene
product or products are introduced into the cell simultaneously or separately[;
and

the transgenic animal comprises a minichromosome].

44. (Twice Amended) A method of producing a transgenic animal,
comprising:

introducing a [DNA] nucleic acid fragment into a cell, wherein the
[DNA] nucleic acid fragment comprises a selectable marker;

growing the cell under selective conditions to produce cells that
have incorporated the [DNA] nucleic acid fragment into their genomic DNA;

selecting a cell that comprises a minichromosome that is about
10 Mb to about 50 Mb that comprises a neocentromere, the selectable marker
and euchromatin;

introducing into the cell DNA encoding a gene product or products;

growing the cell under selective conditions, whereby cells
comprising minichromosomes comprising the DNA encoding the gene product(s)
are produced; [and]

isolating the minichromosome and introducing it into an animal cell;

introducing the cell comprising the minichromosome into a female animal;
and

allowing the cell introduced into the female animal to develop into a
transgenic animal comprising a minichromosome.

B⁶ sub 63) 65. (Amended) The method of claim [64]32, wherein the animal cell is a fertilized ovum.

B⁷ 67. (Amended) The method of claim [64]32, wherein the satellite artificial chromosome is a megachromosome derived from a cell line having all of the identifying characteristics of the cell line deposited under ECACC accession number 96040928 or 96040929.

sub D4) 73. (Amended) A method for producing a transgenic animal, comprising
introducing DNA encoding a gene product or products into a cell
containing the minichromosome of cell line EC3/7C5;
growing the cell under selective conditions, whereby cells
comprising minichromosomes comprising the DNA encoding the gene product(s)
are produced;
isolating the minichromosome and introducing it into an animal cell;
introducing the cell comprising the minichromosome into a female
animal; and
allowing the cell introduced into the female animal to develop into a
transgenic animal comprising a minichromosome [and
exposing the animal cell containing the minichromosome to
condition whereby a transgenic animal develops therefrom].

B⁸ 74. (Amended) A method for producing a transgenic animal, comprising
introducing DNA encoding a gene product or products into a cell containing the
 λ neo-chromosome of cell line KE1 2/4;
growing the cell under selective conditions, whereby cells comprising the
 λ neo-chromosome comprising the DNA encoding the gene product(s) are
produced;
isolating the λ neo-chromosome and introducing it into an animal cell;
introducing the cell comprising the minichromosome into a female animal;
and